Filed: 10/28/02

Applicant: Stefan Kwiatkowski et al.
Title: Dichlorinated Heterocyclic

Compounds And Methods Of Synthesis

Atty. Docket No.: CUTLER-08522 Sheet 1 of 7

$$\begin{array}{c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & \\ & & \\ & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & &$$

$$O = \left\langle \begin{array}{c} \mathbf{F} \\ \mathbf{S} - \mathbf{F} \\ \mathbf{O} \end{array} \right\rangle$$

			Title: Dichlorinated Heterocyclic Compounds And Methods Of Synthesis											
ſ	r	ı		ŀ	Atty. [Docke	t No.:	CUI	ΓLER-	08522	2 Sh	eet 2	of 7	\ 1
!	₹_ ∓													
•	₹_													
			_									Σ		
١.	<u>ာ</u>		127 µM									93.2 µM		l
	<u> </u>		127									93		
┢	8	_												
	FION 50 100				[2						[2]		\square	8
	Ε Ω	-							_	[2]				
	% INHIBITION 3 -50 0 50 1	-	13	КА		.131			_1 \		<u> </u>	1		
	<u>№</u> %	-												
-	ر -100	-												
	7	%	22	45	83	49	56	œ	9	27	62	61	71	64
			+				Σ	Σ	Σ	Ę	Σ	Σ	Σ	
١	2		100 µМ	1000 µM	호	으	잃	ユ 0	Mu e	100 µM	д Д	Mu 000	300 µM	100 µM
	CONC.		우	100	8	9	(r)	_		P	7	5	ĕ	7
	٣		8	8	01	0	8	8	8	8	8	0	0	8
	٥.									5	5	5		
	SP		BOV	167 BOV						₩ E	HOM	MOH		
	÷		70	7										
	BATCH*SPP.		4397	4416						43971	43972	44169		
	Ω		4	4						4	4	4		
			四							E 2	E3			
١			18							8	9			
			SE							SE	SE			
			₹							8	Æ			
			12							STE	STE	 - 		
	Light State of the									Ä	Ä			
	TARGET			! !						호	Ö			
	1		190							SP.	dS	· i		
			۱ğ							Ÿ	Ç			
			146000 PHOSPHODIESTERASE PDE1	•						148000 PHOSPHODIESTERASE PDE2	152000 PHOSPHODIESTERASE PDE3	•		
	#.									000				
	CAT.#		146)						148	15. C	2		
	_		'		_					•	_			

Applicant: Stefan Kwiatkowski et al.

Filed: 10/28/02

*BATCH: REPRESENTS COMPOUNDS TESTED CONCURRENTLY IN THE SAME ASSAY(S).

◆ DENOTES ITEM MEETING THE CRITERIA FOR SIGNIFICANCE

(NEGATIVE VALUES CORRESPOND TO STIMULATION OF BINDING OR ENZYME ACTIVITY) RESULTS WITH≥ 50% STIMULATION OR INHIBITION ARE BOLDFACED.

R = ADDITIONAL COMMENTS BOV = BOVINE; HUM = HUMAN FIG. 2A

				Appli Title:		Stefa Dichlo Comp Synth	orinat ound	ted H	etero	cyclic	Of		
<u>د</u>				Atty.	Dock	et No	.: CL	JTLEI	R-085	22 S	heet 3	s of 7	4
n H								•					
¥													
IC ₅₀							127 µM						
% INHIBITION 1-50 0 50 100	*	Ø	[2]		[3]					8	-		[2]
NE OF	→			<u> </u>	<u> </u>	_KY_	.KL	IXI	N		121		
100 -50	-	-											•
T	%	36	22	8	32	69	20	9/	09	32	48	9	36
CONC.		30 µM	10 µM	Э дМ	100 µM	100 µM	1000 µM	300 µM	100 µM	30 µM	10 µM	Mul 6	100 µM
٤		7	8	0	0	0	7	0	0	8	0	8	7
SPP.					HOM	HOM	M						%
BATCH*SPP.					44171 F	43974 F	44172 H						43975 BOV
					ERASE PDE4	ERASE PDE5							ERASE PDE6
TARGET					154000 PHOSPHODIESTERASE PDE4	156000 PHOSPHODIESTERASE PDE5							156100 PHOSPHODIESTERASE PDE6
CAT.#					154000	156000							156100

Applicant: Stefan Kwiatkowski et al.

Filed: 10/28/02

*BATCH: REPRESENTS COMPOUNDS TESTED CONCURRENTLY IN THE SAME ASSAY(S).

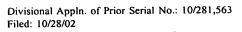
DENOTES ITEM MEETING THE CRITERIA FOR SIGNIFICANCE

(NEGATIVE VALUES CORRESPOND TO STIMULATION OF BINDING OR ENZYME ACTIVITY) RESULTS WITH≥ 50% STIMULATION OR INHIBITION ARE BOLDFACED.

R = ADDITIONAL COMMENTS

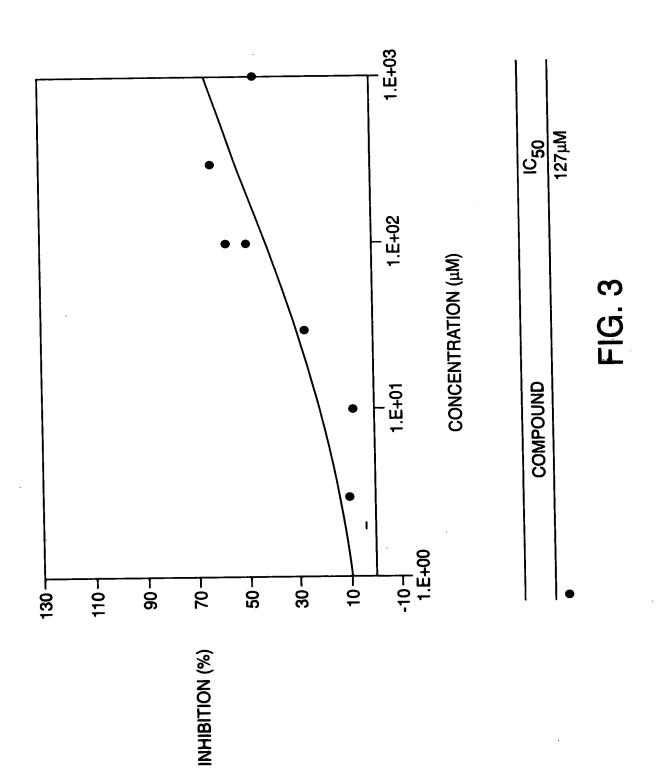
BOV = BOVINE; HUM = HUMAN

FIG. 2B



Applicant: Stefan Kwiatkowski et al.
Title: Dichlorinated Heterocyclic Compounds And Methods Of Synthesis

Atty. Docket No.: CUTLER-08522 Sheet 4 of 7

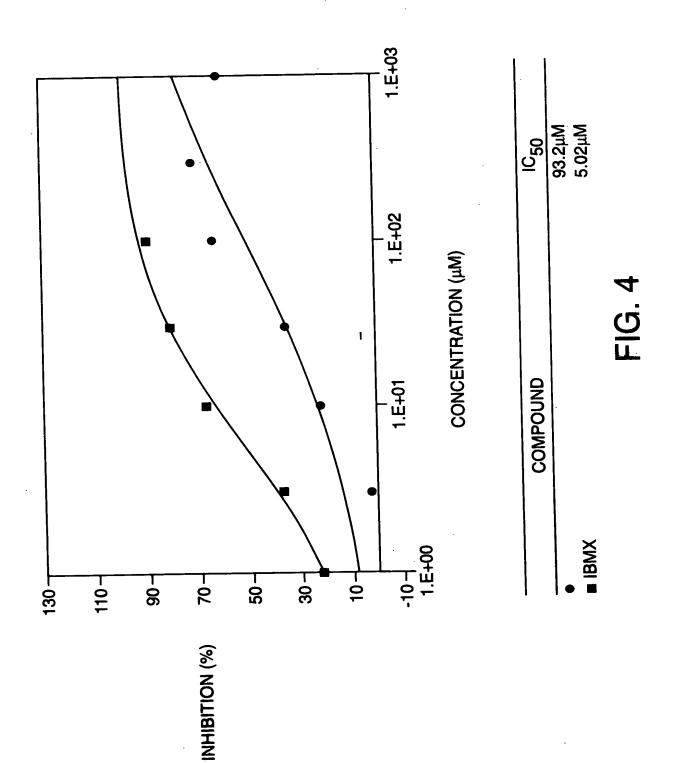


Divisional Appln. of Prior Serial No.: 10/281,563 Filed: 10/28/02

Applicant: Stefan Kwiatkowski et al.
Title: Dichlorinated Heterocyclic Compounds And Methods Of

Synthesis

Atty. Docket No.: CUTLER-08522 Sheet 5 of 7

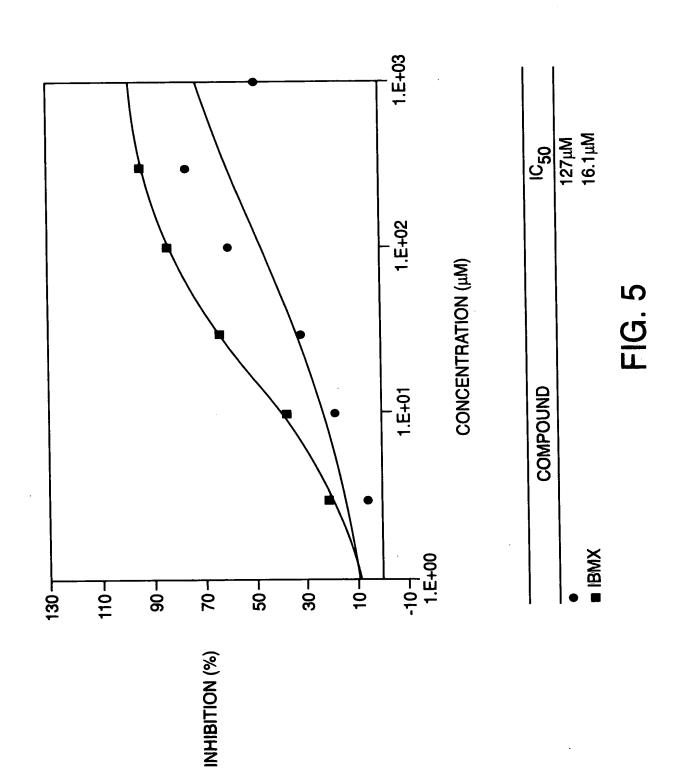


Filed: 10/28/02

Applicant: Stefan Kwiatkowski et al. Dichlorinated Heterocyclic Title: Compounds And Methods Of

Synthesis

Atty. Docket No.: CUTLER-08522 Sheet 6 of 7



Filed: 10/28/02

Applicant: Stefan Kwiatkowski et al.

Title: Dichlorinated Heterocyclic
Compounds And Methods Of

Synthesis

Atty. Docket No.: CUTLER-08522 Sheet 7 of 7